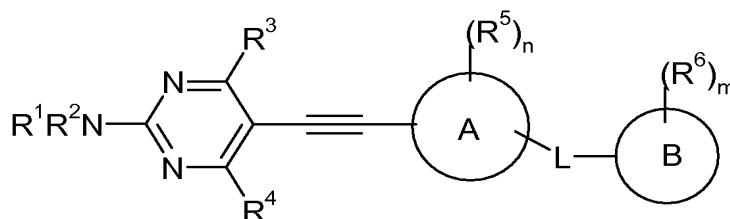


### In the Claims

The listing of claims will replace all prior versions and listings of claims in the application.

### Listings of claims

1. (original): A compound of the Formula I:



Formula I

wherein:

**R<sup>1</sup> and R<sup>2</sup>** are independently selected from hydrogen, (1-6C)alkylsulfonyl, phenyl(CH<sub>2</sub>)<sub>u</sub>- wherein u is 0, 1, 2, 3, 4, 5 or 6, (1-6C)alkanoyl, (1-6C)alkyl, (1-6C)alkoxycarbonyl, (3-6C)cycloalkyl(CH<sub>2</sub>)<sub>x</sub>- in which x is 0, 1, 2, 3, 4, 5 or 6, or a 5 or 6 membered heteroaryl ring, or **R<sup>1</sup> and R<sup>2</sup>** together with the nitrogen atom to which they are attached represent a saturated or partially saturated 3 to 7 membered heterocyclic ring optionally containing another hetero atom selected from N or O;

wherein the (1-6C)alkyl, the (1-6C)alkanoyl and the (3-6C)cycloalkyl groups are optionally substituted by one or more groups independently selected from fluoro, hydroxy, (1-6C)alkyl, (1-6C)alkoxy, (1-6C)alkoxy(1-6C)alkoxy, (1-6C)alkoxy(1-6C)alkoxy(1-6C)alkoxy, amino, mono(1-6C)alkylamino, di-[(1-6C)alkyl]amino, carbamoyl, mono(1-6C)alkylcarbamoyl, di-[(1-6C)alkyl]carbamoyl, -N(R<sup>d</sup>)C(O)(1-6C)alkyl in which R<sup>d</sup> is hydrogen or (1-6C)alkyl, a saturated or partially saturated 3 to 7 membered heterocyclic ring, or a 5 or 6 membered heteroaryl ring,

wherein the (1-6C)alkoxy, (1-6C)alkoxy(1-6C)alkoxy and (1-6C)alkoxy(1-6C)alkoxy(1-6C)alkoxy groups and the (1-6C)alkyl groups of the mono(1-6C)alkylamino, di-[(1-6C)alkyl]amino, mono(1-6C)alkylcarbamoyl, di-[(1-6C)alkyl]carbamoyl and/or -N(R<sup>d</sup>)C(O)(1-6C)alkyl groups are optionally substituted by one or more hydroxy groups;

wherein the phenyl is optionally substituted by one or more groups independently selected from halo, (1-6C)alkyl, (1-6C)alkoxy, amino, mono(1-6C)alkylamino or di-[(1-6C)alkyl]amino, wherein the (1-6C)alkyl and the (1-6C)alkoxy groups

are optionally substituted by one or more groups independently selected from hydroxy, amino, mono(1-6C)alkylamino or di-[(1-6C)alkyl]amino; and wherein any heterocyclic and heteroaryl rings within  $R^1$  and/or  $R^2$  are optionally independently substituted by one or more of the following: (1-4C)alkyl, (1-4C)alkoxy, (1-4C)alkoxy(1-4C)alkyl, hydroxy, amino, mono(1-6C)alkylamino, di-[(1-6C)alkyl]amino, a saturated or partially saturated 3 to 7 membered heterocyclic ring or  $-C(O)(CH_2)_zY$  wherein  $z$  is 0, 1, 2 or 3 and  $Y$  is selected from hydrogen, hydroxy, (1-4C)alkoxy, amino, mono(1-6C)alkylamino, di-[(1-6C)alkyl]amino or a saturated or partially saturated 3 to 7 membered heterocyclic ring; and provided that when  $R^1$  and/or  $R^2$  is a (1C)alkanoyl group, then the (1C)alkanoyl is not substituted by fluoro or hydroxy;

**$R^3$  and  $R^4$**  are independently selected from hydrogen, (1-6C)alkyl or (1-6C)alkoxy, wherein the (1-6C)alkyl and the (1-6C)alkoxy groups are optionally substituted by one or more groups independently selected from: fluoro, hydroxy, (1-6C)alkyl, (1-6C)alkoxy, amino, mono(1-6C)alkylamino, di-[(1-6C)alkyl]amino, carbamoyl, mono(1-6C)alkylcarbamoyl or di-[(1-6C)alkyl]carbamoyl, a saturated or partially saturated 3 to 7 membered heterocyclic ring or a 5 or 6 membered heteroaryl ring, wherein said heterocyclic and heteroaryl rings are optionally independently substituted by one or more of the following: (1-4C)alkyl, (1-4C)alkoxy, hydroxy, amino, mono(1-6C)alkylamino or di-[(1-6C)alkyl]amino or a saturated or partially saturated 3 to 7 membered heterocyclic ring; or one of  **$R^3$**  and  **$R^4$**  is as defined above and the other represents a group  $-NR^1R^2$  as defined above;

**A** represents an aryl group or a 5 or 6 membered heteroaryl ring selected from furyl, pyrrolyl, thienyl, oxazolyl, isoxazolyl, imidazolyl, pyrazolyl, thiazolyl, isothiazolyl, oxadiazolyl, thiadiazolyl, triazolyl, tetrazolyl, pyridyl, pyridazinyl, pyrimidinyl, pyrazinyl or 1,3,5-triazinyl;

**$R^5$**  is selected from cyclopropyl, cyano, halo, (1-6C)alkoxy or (1-6C)alkyl, wherein the (1-6C)alkyl and the (1-6C)alkoxy groups are optionally substituted by cyano or by one or more fluoro;

**n** is 0, 1, 2 or 3;

**L** is attached meta or para on ring A with respect to the point of attachment of the ethynyl group and represents  $-C(R^aR^b)C(O)N(R^9)-$ ,  $-N(R^8)C(O)C(R^aR^b)-$ ,  $-N(R^8)C(O)N(R^9)-$ ,  $-N(R^8)C(O)O-$ , or  $-OC(O)-N(R^9)-$ , wherein  $R^8$  and  $R^9$  independently represent hydrogen or (1-6C)alkyl and wherein  $R^a$  and  $R^b$  independently represent hydrogen or (1-6C)alkyl or  $R^a$  and  $R^b$  together with the carbon atom to which they are attached represent (3-6C)cycloalkyl;

**B** represents a (3-7C)cycloalkyl ring, a saturated or partially saturated 3 to 7 membered heterocyclic ring, an aryl group, a 5 or 6 membered heteroaryl ring selected from furyl, pyrrolyl, thienyl, oxazolyl, isoxazolyl, imidazolyl, pyrazolyl, thiazolyl, isothiazolyl, oxadiazolyl, thiadiazolyl, triazolyl, tetrazolyl, pyridyl, pyridazinyl, pyrimidinyl, pyrazinyl or 1,3,5-triazinyl, or a 8, 9 or 10 membered bicyclic group which optionally contains 1, 2, 3 or 4 heteroatoms independently selected from N, O and S and which is saturated, partially saturated or aromatic;

**R<sup>6</sup>** is selected from halo, cyano, oxo, a (3-7C)cycloalkyl ring, a saturated or partially saturated 3 to 7 membered heterocyclic ring, and  $-N(R^c)C(O)(1-6C)alkyl$  in which  $R^c$  is hydrogen or (1-6C)alkyl; or

**R<sup>6</sup>** is selected from (1-6C)alkyl,  $-S(O)_p(1-6C)alkyl$  wherein p is 0, 1 or 2, or (1-6C)alkoxy, wherein the (1-6C)alkyl,  $-S(O)_p(1-6C)alkyl$  and the (1-6C)alkoxy groups are optionally substituted by one or more groups independently selected from cyano, fluoro, hydroxy, (1-6C)alkoxy, amino, mono(1-6C)alkylamino, di-[(1-6C)alkyl]amino, a (3-7C)cycloalkyl ring or a saturated or partially saturated 3 to 7 membered heterocyclic ring; and

wherein the (3-7C)cycloalkyl ring and saturated or partially saturated 3 to 7 membered heterocyclic ring are optionally independently substituted by one or more groups selected from (1-6C)alkyl; and

**m** is 0, 1, 2 or 3;

and when **B** is a (3-7C)cycloalkyl ring, a saturated or partially saturated 3 to 7 membered heterocyclic ring or a saturated or partially saturated 8, 9 or 10 membered bicyclic group, the rings and the bicyclic group optionally bear 1 or 2 oxo or thioxo substituents;

and salts thereof.

2. (original): A compound of Formula I according to Claim 1, wherein:

**R<sup>6</sup>** is selected from halo, cyano, a (3-7C)cycloalkyl ring, a saturated or partially saturated 3 to 7 membered heterocyclic ring or an alkanoylamino group -N(R<sup>c</sup>)C(O)(1-6C)alkyl in which R<sup>c</sup> is hydrogen or (1-6C)alkyl; or

**R<sup>6</sup>** is selected from (1-6C)alkyl or (1-6C)alkoxy, wherein the (1-6C)alkyl and the (1-6C)alkoxy groups are optionally substituted by one or more groups independently selected from cyano, fluoro, hydroxy, (1-6C)alkoxy, amino, mono(1-6C)alkylamino, di-[(1-6C)alkyl]amino, a (3-7C)cycloalkyl ring or a saturated or partially saturated 3 to 7 membered heterocyclic ring;

and salts thereof.

3. (original): A compound of the Formula I according to claim 1, wherein:

**R<sup>1</sup> and R<sup>2</sup>** are independently selected from hydrogen, (1-6C)alkylsulfonyl, phenyl(CH<sub>2</sub>)<sub>u</sub>- wherein u is 0, 1, 2, 3, 4, 5 or 6, (1-6C)alkanoyl, (1-6C)alkyl, (1-6C)alkoxycarbonyl, or (3-6C)cycloalkyl(CH<sub>2</sub>)<sub>x</sub>- in which x is 0, 1, 2, 3, 4, 5 or 6 or **R<sup>1</sup> and R<sup>2</sup>** together with the nitrogen atom to which they are attached represent a saturated or partially saturated 3 to 7 membered heterocyclic ring optionally containing another hetero atom selected from N or O;

wherein the alkyl and the cycloalkyl groups are optionally substituted by one or more groups selected from fluoro, hydroxy, (1-6C)alkyl, (1-6C)alkoxy, amino, mono(1-6C)alkylamino or di-[(1-6C)alkyl]amino, a saturated or partially saturated 3 to 7 membered heterocyclic ring or a 5 or 6 membered heteroaryl ring, wherein said heterocyclic and heteroaryl rings are optionally independently substituted by one or more of the following: (1-4C)alkyl, hydroxy, amino, mono(1-6C)alkylamino or di-[(1-6C)alkyl]amino or a saturated or partially saturated 3 to 7 membered heterocyclic ring;

and wherein the phenyl is optionally substituted by one or more groups selected from halo, (1-6C)alkyl, (1-6C)alkoxy, amino, mono(1-6C)alkylamino or di-[(1-6C)alkyl]amino, wherein the (1-6C)alkyl or (1-6C)alkoxy are optionally substituted by hydroxy, amino, mono(1-6C)alkylamino or di-[(1-6C)alkyl]amino;

**R<sup>3</sup> and R<sup>4</sup>** are independently selected from hydrogen, (1-6C)alkyl or (1-6C)alkoxy wherein the alkyl and the alkoxy groups are optionally substituted by one or more

groups selected from fluoro, hydroxy, (1-6C)alkyl, (1-6C)alkoxy, amino, mono(1-6C)alkylamino or di-[(1-6C)alkyl]amino, a saturated or partially saturated 3 to 7 membered heterocyclic ring or a 5 or 6 membered heteroaryl ring, wherein said heterocyclic and heteroaryl rings are optionally independently substituted by one or more of the following: (1-4C)alkyl, hydroxy, amino, mono(1-6C)alkylamino or di-[(1-6C)alkyl]amino or a saturated or partially saturated 3 to 7 membered heterocyclic ring;

or one of  $R^3$  and  $R^4$  is as defined above and the other represents a group  $-NR^1R^2$  as defined above;

$R^5$  is selected from cyano, halo, (1-6C)alkoxy or (1-6C)alkyl optionally substituted by cyano or by one or more fluoro;

**B** represents a (3-7C)cycloalkyl ring, an aryl or a 5 or 6 membered heteroaryl ring selected from furyl, pyrrolyl, thienyl, oxazolyl, isoxazolyl, imidazolyl, pyrazolyl, thiazolyl, isothiazolyl, oxadiazolyl, thiadiazolyl, triazolyl, tetrazolyl, pyridyl, pyridazinyl, pyrimidinyl, pyrazinyl or 1,3,5-triazinyl;

$R^6$  is selected from halo, cyano, a saturated or partially saturated 3 to 7 membered heterocyclic ring or an alkanoylamino group  $-N(R^c)C(O)(1-6C)alkyl$  in which  $R^c$  is hydrogen or (1-6C)alkyl; or  $R^6$  is selected from (1-6C)alkyl or (1-6C)alkoxy, wherein the alkyl and the alkoxy groups are optionally substituted by one or more groups selected from cyano, fluoro, hydroxy, (1-6C)alkoxy, amino, mono(1-6C)alkylamino, di-[(1-6C)alkyl]amino, or a saturated or partially saturated 3 to 7 membered heterocyclic ring; and

m is **0, 1, 2 or 3**; and when m is at least 2 then two substituents on adjacent carbon atoms in ring B may together represent a methylenedioxy group;

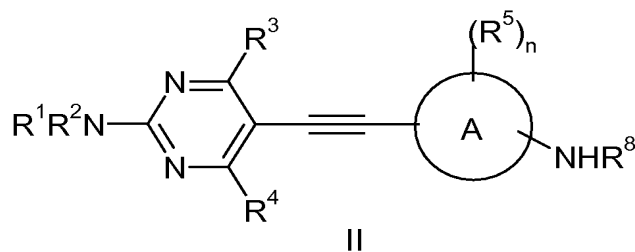
and wherein A, L and n are as defined in Claim 1.

and salts thereof.

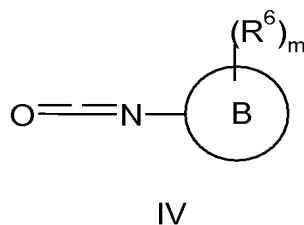
4. (currently amended): A compound according to ~~any one of Claims 1, 2 and 3~~ wherein A is selected from phenyl, pyridyl, thiazolyl, thiadiazolyl or pyrimidinyl.

5. (currently amended): A compound accordingly to ~~any one of the preceding claims~~ claim 1 wherein B is selected from phenyl, 2,3-di-hydro-indenyl, piperidinyl, pyridyl, pyrazolyl, isothiazolyl, thiadiazolyl, isoxazolyl, benzodioxinyl, benzodioxolyl or tetrahydropyranyl
6. (currently amended): A compound accordingly to ~~any one of the preceding claims~~ claim 1 wherein L is selected from  $-N(R^8)C(O)N(R^9)-$ ,  $-N(R^8)C(O)O-$  or  $-N(R^8)C(O)CH_2-$  wherein  $R^8$  and  $R^9$  independently represent hydrogen or (1-6C)alkyl.
7. (currently amended): A compound accordingly to ~~any one of the preceding claims~~ claim 1 wherein  $R^1$  and  $R^2$  are both hydrogen or  $R^1$  is hydrogen or (1-6C)alkyl and  $R^2$  is (1-6C)alkyl  
wherein (1-6Calkyl) is optionally substituted by hydroxy, amino, mono(1-6C)alkylamino or di(1-6C)alkylamino, carbamoyl, (1-6C)alkoxy, (1-6C)alkoxy(1-6C)alkoxy,  $-N(R^d)C(O)(1-6C)alkyl$  in which  $R^d$  is hydrogen or (1-6C)alkyl, aryl (particularly phenyl), a saturated or partially saturated 3 to 7 membered heterocyclic ring or a 5 or 6 membered heteroaryl ring;  
wherein the (1-6C)alkoxy, mono(1-6C)alkylamino and  $-N(R^d)C(O)(1-6C)alkyl$  groups are optionally substituted by hydroxy;  
wherein an aryl ring, a saturated or partially saturated 3 to 7 membered heterocyclic ring or a 5 or 6 membered heteroaryl ring is optionally substituted by (1-4C)alkyl, (1-4C)alkoxy or  $-C(O)CH_2Y$  wherein Y is selected from hydroxy or di(1-6C)alkylamino.
8. (currently amended): A compound accordingly to ~~any one of the preceding claims~~ claim 1 wherein  $R^3$  and  $R^4$  are both hydrogen.
9. (currently amended): A compound accordingly to ~~any one of the preceding claims~~ claim 1 wherein  $R^6$  is independently selected from halo, cyano, oxo, (3-7C)cycloalkyl, a saturated 3 to 7 membered heterocyclic ring (optionally substituted by (1-4C)alkyl),  $-N(R^c)C(O)(1-6C)alkyl$  wherein  $R^c$  is hydrogen or (1-6C)alkyl (particularly (1-4C)alkyl), (1-6C)alkyl (optionally substituted by up to three groups independently selected from halo) or (1-6C)alkoxy and m is selected from 1 or 2.
10. (original): A compound according to Claim 1 which is any one or more of examples 1 to 152 or a salt thereof.

11. (currently amended): A pharmaceutical composition which comprises a compound of the Formula I, or a pharmaceutically acceptable salt thereof, as defined in claims 1 to 40 in association with a pharmaceutically acceptable diluent or carrier.
12. (canceled)
13. (canceled)
14. (canceled)
15. (currently amended): A process for preparing a compound of formula I, or salt thereof, as defined in Claim 1, or a pharmaceutically acceptable salt thereof (wherein  $R^1$ ,  $R^2$ ,  $R^3$ ,  $R^4$ ,  $R^5$ ,  $R^6$ ,  $R^7$ ,  $R^8$ ,  $R^9$ ,  $R^{10}$ ,  $R^{11}$ , L, ring A and ring B, n and m are, unless otherwise specified, as defined in Claim 1) comprising:
  - (a) For compounds of the formula I wherein L is  $-N(R^8)C(O)N(H)-$ , the reaction of a compound of the formula II:

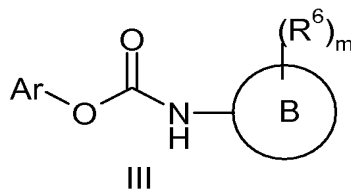


wherein  $R^1$ ,  $R^2$ ,  $R^3$ ,  $R^4$ ,  $R^5$ ,  $R^8$ , n and A have any of the meanings defined hereinbefore except that any functional group is protected if necessary, with an isocyanate of the formula IV:



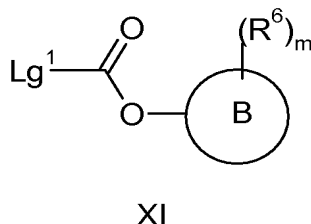
wherein  $R^6$ , m and B have any of the meanings defined hereinbefore except that any functional group is protected if necessary; or

- (b) For compounds of the formula I wherein L is  $-N(R^8)C(O)N(H)-$ , the reaction of a compound of the formula II as defined above with an aryl carbamate of the formula III:



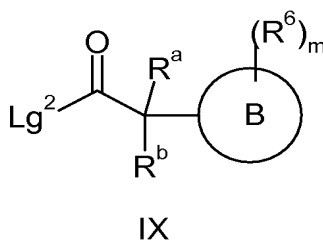
wherein Ar is a suitable aryl group, ~~for example phenyl~~, and  $R^6$ , m and B have any of the meanings defined hereinbefore except that any functional group is protected if necessary; or

- (c) For compounds of the formula I wherein L is  $N(R^8)C(O)-O-$ , the reaction of a compound of the formula II as defined above with a compound of the formula XI:



wherein  $Lg^1$  is a suitable displaceable group ~~for example halogeno (such as fluoro, chloro or bromo)~~ and  $R^6$ , m and B have any of the meanings defined hereinbefore except that any functional group is protected if necessary; or

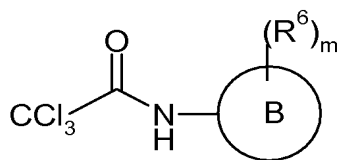
- (d) For compounds of the formula I wherein L is  $N(R^8)C(O)C(R^aR^b)$ , the reaction of a compound of the formula II as defined above with a compound of the formula IX:



wherein  $Lg^2$  is a suitable displaceable group,  $R^x-C(O)-O-$  or  $R^x-O-$  (wherein  $R^x$  is a suitable alkyl or aryl group) and  $R^6$ ,  $R^a$ ,  $R^b$ , m and B have any of the meanings defined hereinbefore except that any functional group is protected if necessary; or

- (e) For compounds of the formula I wherein L is  $-N(R^8)C(O)N(H)-$ , the reaction of a compound of the formula II as defined above with a trichloroacetylamine of the formula XIII:

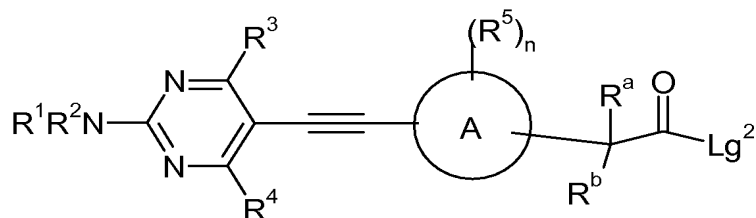




XIII

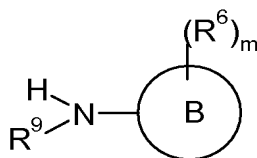
wherein  $R^6$ ,  $m$  and  $B$  have any of the meanings defined hereinbefore except that any functional group is protected if necessary; or

- (f) For compounds of the formula I wherein  $L$  is  $-C(R^a R^b)C(O)N(R^9)-$ , the reaction of a compound of the formula XIV:



XIV

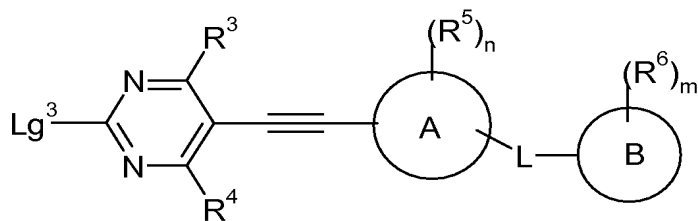
wherein  $Lg^2$  is a suitable displaceable group as described above and  $R^1$ ,  $R^2$ ,  $R^3$ ,  $R^4$ ,  $R^5$ ,  $R^a$ ,  $R^b$ ,  $n$  and  $A$  have any of the meanings defined hereinbefore except that any functional group is protected if necessary, with an amine of the formula XV:



XV

wherein  $R^6$ ,  $R^9$ ,  $m$  and  $B$  have any of the meanings defined hereinbefore except that any functional group is protected if necessary; or

- (g) The reaction of a compound of the formula XVI:

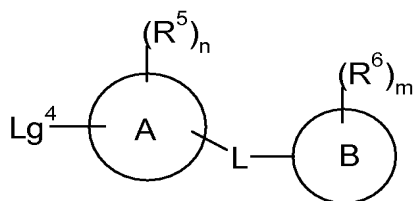


XVI

wherein  $Lg^3$  is a suitable displaceable group for example halogeno (such as fluoro, chloro, bromo or iodo), methyl sulfonyl, methylthio or aryloxy (such as phenoxy) and  $R^3$ ,  $R^4$ ,  $R^5$ ,  $R^6$ ,  $n$ ,  $m$ ,  $A$ ,  $B$  and  $L$  have any of the meanings defined

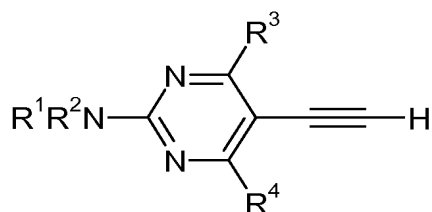
hereinbefore except that any functional group is protected if necessary, with an amine of the formula  $\text{HNR}^1\text{R}^2$ , wherein  $\text{R}^1$  and  $\text{R}^2$  have any of the meanings defined hereinbefore except that any functional group is protected if necessary; or

- (h) The reaction of a compound of the formula XVII:



XVII

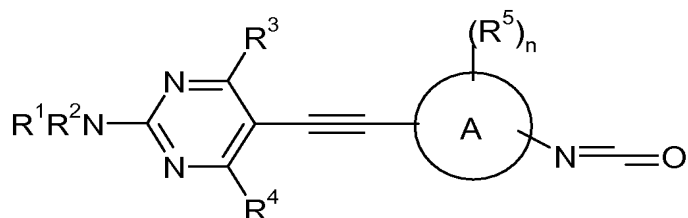
wherein  $\text{Lg}^4$  is a suitable displaceable group ~~for example halogeno (such as chloro, bromo or iodo) or a sulfonyloxy group (such as trifluoromethylsulfonyloxy)~~ and  $\text{R}^5$ ,  $\text{R}^6$ ,  $n$ ,  $m$ ,  $A$ ,  $B$  and  $L$  have any of the meanings defined hereinbefore except that any functional group is protected if necessary, with an alkyne of the formula XVIII:



XVIII

wherein  $\text{R}^1$ ,  $\text{R}^2$ ,  $\text{R}^3$  and  $\text{R}^4$  have any of the meanings defined hereinbefore except that any functional group is protected if necessary; or

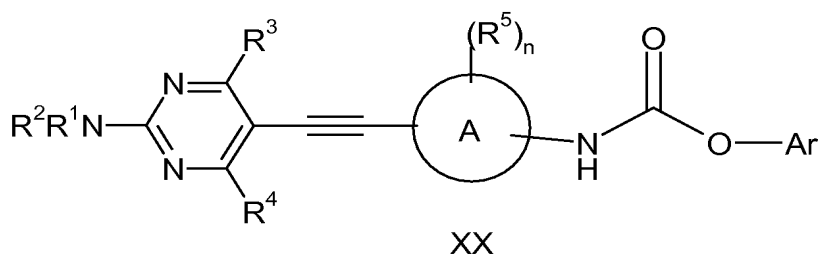
- (i) For compounds of the formula I wherein  $L$  is  $-\text{N}(\text{H})\text{C}(\text{O})\text{N}(\text{R}^9)-$ , the reaction of an isocyanate of the formula XIX:



XIX

wherein  $\text{R}^1$ ,  $\text{R}^2$ ,  $\text{R}^3$ ,  $\text{R}^4$ ,  $\text{R}^5$ ,  $n$  and  $A$  have any of the meanings defined hereinbefore except that any functional group is protected if necessary, with an amine of the formula XV as defined above; or

- (j) For compounds of the formula I wherein  $L$  is  $-\text{N}(\text{H})\text{C}(\text{O})\text{N}(\text{R}^9)-$ , the reaction of a compound of the formula XX:



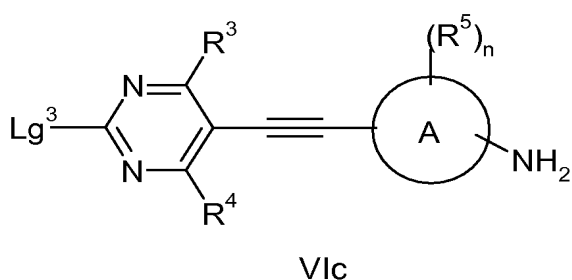
wherein Ar is a suitable aryl group, ~~for example phenyl~~, and  $R^1$ ,  $R^2$ ,  $R^3$ ,  $R^4$ ,  $R^5$ ,  $n$  and A have any of the meanings defined hereinbefore except that any functional group is protected if necessary, with an amine of the formula XV as defined above.

and thereafter if necessary:

- i) converting a compound of the Formula (I) into another compound of the Formula (I);
- ii) removing any protecting groups;
- iii) forming a salt.

16. (currently amended): A compound selected from Formulae II, XIV, XVI, XIX and XX as defined in Claim 15, wherein A is a 5 or 6 membered heteroaryl ring selected from furyl, pyrrolyl, thienyl, oxazolyl, isoxazolyl, imidazolyl, pyrazolyl, thiazolyl, isothiazolyl, oxadiazolyl, thiadiazolyl, triazolyl, tetrazolyl, pyridyl, pyridazinyl, pyrimidinyl, pyrazinyl or 1,3,5-triazinyl

or a compound of Formula VIc:



or salt thereof, wherein A is a 5 or 6 membered heteroaryl ring selected from furyl, pyrrolyl, thienyl, oxazolyl, isoxazolyl, imidazolyl, pyrazolyl, thiazolyl, isothiazolyl, oxadiazolyl, thiadiazolyl, triazolyl, tetrazolyl, pyridyl, pyridazinyl, pyrimidinyl, pyrazinyl or 1,3,5-triazinyl and  $Lg^3$ ,  $R^3$ ,  $R^4$ ,  $R^5$  and  $n$  are as defined in Claim 15.

17. (new): A method of inhibiting Tie2 receptor tyrosine kinase in a warm-blooded animal in need of such treatment, which comprises administering to said animal an effective amount of a compound of the formula I, or a pharmaceutically acceptable salt thereof, according to claim 1.

18. (new): A method for producing an anti-angiogenic effect in a warm-blooded animal in need of such treatment, which comprises administering to said animal an effective amount of a compound of the formula I, or a pharmaceutically acceptable salt thereof, as claimed in claim 1.